Appointment

From: Pinto, Ana [Pinto.Ana@epa.gov]
Sent: 3/1/2021 12:41:20 PM
To: Rick Reiss [rreiss@exponent.com]; Perron, Monique [Perron.Monique@epa.gov]; Lowit, Anna [Lowit.Anna@epa.gov]; Mendez, Elizabeth [Mendez.Elizabeth@epa.gov]; Tan, Cecilia [Tan.Cecilia@epa.gov]; Miller, David [Miller.DavidJ@epa.gov]; Villanueva, Philip [Villanueva.Philip@epa.gov]; Nguyen, James [Nguyen.James@epa.gov]; Padilla, Stephanie [Padilla.Stephanie@epa.gov]; Sherman, Kelly [Sherman.Kelly@epa.gov]; Mannix, Marianne [Mannix.Marianne@epa.gov]; Jakob, Avivah [Jakob.Avivah@epa.gov]
CC: csmith@gowanco.com; Betsy Codrea [BCodrea@GOWANCO.com]; Paul Whatling [Paul.Whatling@fmc.com]; Christian Strupp [CStrupp@gowanco.com]

Subject: SAP Report Follow-Up Meeting with Exponent
Attachments: Meek et al.pdf; Herkert 2012.pdf
Location: Microsoft Teams Meeting

Start: 3/3/2021 7:00:00 PM
End: 3/3/2021 8:00:00 PM
Show Time As: Tentative

Required Attendees: Rick Reiss; Perron, Monique; Anna Lowitt (Lowit.Anna@epa.gov); Mendez, Elizabeth; Tan, Cecilia; Miller, David; Villanueva, Philip; Nguyen, James; Padilla, Stephanie; Sherman, Kelly; Mannix, Marianne; Jakob, Avivah
Optional Attendees: Cindy Smith; Betsy Codrea; Paul Whatling; Christian Strupp

Meeting between Exponent and EPA/ORD to discuss next steps now that the SAP report has been released.

Proposed Agenda for OP In Vitro Call:

1. Age sensitivity of rats. Questions were raised about our exclusive use of adult rats and whether we needed to also have data for juvenile rats. New information is available from the attached study by Meek et al. (Jan Chambers lab) that shows comparable IC50s for 12 OPs (no overlap with our study) for brain AChE derived from PND1, PND11, and PND70 rats (see Table 5).
2. Use of AChE from RBC vs. Brain. Attached paper by Herkert et al. shows brain and RBC AChE inhibition kinetics were "highly comparable."
3. Rudy Richardson piece responding to analysis by Stephanie Padilla. Are further discussions possible on issues on lack of AChE variability among human and the role that biological reasoning (beyond just experiments and statistics) can play in the decisions?
4. IC50 values vs. bimolecular rate constant. Analysis for a limited number of chemicals shows that IC50s have lower variability than bimolecular rate constants. Does this effect the sample size discussion?
5. Adequacy of sample size. Discussion on SAP report. Possible ways to address SAP concerns.
6. Supplemental variability study. Does the supplemental variability study advance the discussion, particularly combined with biological reasoning?
7. Other issues

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